

(FILE 'HOME' ENTERED AT 12:16:23 ON 03 NOV 2005)

FILE 'AGRICOLA, MEDLINE, CAPLUS, BIOSIS' ENTERED AT 12:16:26 ON 03 NOV 2005

L1	11349 S (NEUTRAL (1N) ENDOPEPTIDASE) OR NEP OR NEPRILYSIN
L2	900 S L1 AND (MUS OR MOUSE)
L3	323 S L2 AND (CDNA OR CLON? OR GENE)
L4	105 S L3 AND PY<1999
L5	69 DUP REM L4 (36 DUPLICATES REMOVED)
L6	0 S L5 AND 765

L6 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2005 ACS on STN
 AN 1993:186782 CAPLUS
 DN 118:186782
 TI Murine common acute lymphoblastic leukemia antigen (CD10 neutral endopeptidase 24.11). Molecular characterization, chromosomal localization, and modeling of the active site
 AU Chen, Chang Yan; Salles, Gilles; Seldin, Michael F.; Kister, Alexander E.; Reinherz, Ellis L.; Shipp, Margaret A.
 CS Lab. Immunobiol., Harvard Med. Sch., Boston, MA, 02115, USA
 SO Journal of Immunology (1992), 148(9), 2817-25
 CODEN: JOIMA3; ISSN: 0022-1767
 DT Journal
 LA English
 AB To further analyze antigen CD10/neutral endopeptidase 24.11 CD10/NEP] function in lymphoid and nonlymphoid cells using well characterized murine systems, the murine CD10/NEP homolog was isolated, its chromosomal location was determined and the enzyme active site was modeled. The murine CD10/NEP **cDNA** predicts a 750-amino acid (aa) type II integral membrane protein with 90% identity to the human CD10 sequence and 100% conservation of critical aa and functional motifs. The latter include the pentapeptide consensus sequence required for zinc binding and catalytic activity, addnl. aa associated with substrate binding, and the extracellular cysteines that participate in disulfide bonds required for enzymic activity. Like its human homolog, murine CD10/NEP has multiple alternative 5'-untranslated region sequences. The gene is localized on the proximal half of murine chromosome 3. In Northern anal., murine CD10/NEP transcripts are abundant in bone marrow stromal cells that support pre-B cell differentiation but are undetectable in representative Abelson transformed pre-B cell lines. The murine CD10/NEP active site was modeled by aligning critical conserved CD10/NEP residues with comparable residues in the active site of thermolysin, a bacterial metalloprotease with similar substrate specificity. The model predicts that the 2 enzymes have similar clefts that comprise the active site and permit zinc-dependent substrate interactions.

L6 ANSWER 8 OF 12 MEDLINE on STN DUPLICATE 4
 AN 93390947 MEDLINE
 DN PubMed ID: 8397369
 TI NEP: a novel receptor-like tyrosine kinase expressed in proliferating neuroepithelia.
 AU Zerlin M; Julius M A; Goldfarb M
 CS Department of Biochemistry and Molecular Biophysics, Columbia University College of Physicians and Surgeons, New York, New York 10032.
 SO Oncogene, (1993 Oct) 8 (10) 2731-9.
 Journal code: 8711562. ISSN: 0950-9232.
 CY ENGLAND: United Kingdom
 DT Journal; Article; (JOURNAL ARTICLE)
 LA English
 FS Priority Journals
 EM 199310
 ED Entered STN: 19931105
 Last Updated on STN: 19931105
 Entered Medline: 19931018
 AB We have isolated a murine **cdna**, **nep**, which encodes a novel receptor-like protein tyrosine kinase. The kinase region of NEP protein bears 50% amino acid sequence identity to the neurotrophin receptors (TRKs). While the intracytoplasmic portion of NEP also contains a short kinase insert region and C-terminal tail reminiscent of the TRK proteins, the putative extracellular domain of NEP is unrelated to any known proteins. The **nep** gene is strongly expressed within proliferating neuroepithelia of **mouse** embryos, commencing at the early somite stage (embryonic day 8.0) and persisting in the proliferative ventricular zones of the brain and spinal cord, suggesting that one function of NEP kinase is to signal proliferation of neuroepithelial cells in response to an as yet unknown ligand. The **nep** gene is also expressed in embryonic sensory ganglia, striated muscle and epidermis, as well as in several adult tissues, including the ventricle linings and glia subpopulations in the brain.

Neutral endopeptidase modulation of septic shock.

AU Lu B; Gerard N P; Kolakowski L F Jr; Bozza M; Zurakowski D; Finco O;
Carroll M C; Gerard C
CS Ina Sue Perlmutter Laboratory, Children's Hospital, Boston, Massachusetts,
USA.
NC HL19170 (NHLBI)
HL51366 (NHLBI)
SO Journal of experimental medicine, (1995 Jun 1) 181 (6) 2271-5.
Journal code: 2985109R. ISSN: 0022-1007.
CY United States
DT Journal; Article; (JOURNAL ARTICLE)
LA English
FS Priority Journals
EM 199506
ED Entered STN: 19950707
Last Updated on STN: 19950707
Entered Medline: 19950623
AB Neutral endopeptidase (NEP; EC. 3.4.24.11) is a type 2 cell surface
metalloprotease known by a variety of eponyms, including enkephalinase,
common acute lymphoblastic leukemia antigen, and CD10. Identified
substrates are largely neural or humoral oligopeptide agonists, and the
enzyme functions to terminate signaling by degrading the ligand,
analogously to acetylcholine/acetylcholinesterase. Targeted disruption of
the **NEP** locus in **mice** results in enhanced lethality to
endotoxin shock with a pronounced gene dosage effect. The site(s) of
action appears downstream from release of tumor necrosis factor and
interleukin-1 since NEP-deficient animals demonstrate increased
sensitivity to these mediators as well. This unexpected finding indicates
an important protective role for NEP in septic shock.

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 635 640 645

cta gca gac aac cag aat gtg aac gga ttc agt acc ctc ggg gag aac 2320
 Leu Ala Asp Asn Gln Asn Val Asn Gly Phe Ser Thr Leu Gly Glu Asn
 650 655 660

att gcc gac aac gga ggt gtg cga cag gca tac aag gct tac cta cgg 2368
 Ile Ala Asp Asn Gly Gly Val Arg Gln Ala Tyr Lys Ala Tyr Leu Arg
 665 670 675

tgg ctg gct gat ggc ggc aaa gat cag cga ctg ccg gga ctg aac ctg 2416
 Trp Leu Ala Asp Gly Gly Lys Asp Gln Arg Leu Pro Gly Leu Asn Leu
 680 685 690 695

acc tat gcc cag ctt ttc ttc atc aac tat gcc cag gtg tgg tgt ggg 2464
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 730 735 740

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 760 765

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 <213> Mus musculus

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 35 40 45
 Ser Leu Leu His Phe Ser Trp Asp Glu Arg Thr Val Val Lys Arg Ala
 50 55 60
 Leu Arg Asp Ser Ser Leu Lys Ser Asp Ile Cys Thr Thr Pro Ser Cys
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 Val Ile Ala Ala Ala Arg Ile Leu Glu Asn Met Asp Gln Ser Arg Asn
 85 90 95
 Pro Cys Glu Asn Phe Tyr Gln Tyr Ala Cys Gly Gly Trp Leu Arg His
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 His Val Ile Pro Glu Thr Asn Ser Arg Tyr Ser Val Phe Asp Ile Leu
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 130 135 140
 Ser Gln His Arg Pro Ala Val Glu Lys Ala Lys Thr Leu Tyr Arg Ser
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 Cys Met Asn Gln Ser Val Ile Glu Lys Arg Asp Ser Glu Pro Leu Leu
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 Ser Val Leu Lys Met Val Gly Gly Trp Pro Val Ala Met Asp Lys Trp
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 195 200 205
 Leu Asn Ser Gln Phe Asn Arg Arg Val Leu Ile Asp Leu Phe Ile Trp
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 225 230 235 240
 Thr Leu Gly Met Pro Ser Arg Glu Tyr Tyr Phe Gln Glu Asp Asn Asn
 245 250 255
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 260 265 270
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 275 280 285
 Glu Glu Met Ala Glu Val Leu Glu Leu Glu Thr His Leu Ala Asn Ala
 290 295 300
 Thr Val Pro Gln Glu Lys Arg His Asp Val Thr Ala Leu Tyr His Arg
 305 310 315 320
 Met Asp Leu Met Glu Leu Gln Glu Arg Phe Gly Leu Lys Gly Phe Asn

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Leu	Glu	Asp	Ile	Ile	Asp	Ser	Tyr	Ser	Ala	Arg	Thr	Met	Gln	Asn	Tyr				
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Asn	Leu	Lys	Asn	Asn	Ala	Gln	Arg	Ser	Leu	Lys	Lys	Leu	Arg	Glu	Lys				
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Val	Asp	Gln	Asn	Leu	Trp	Ile	Ile	Gly	Ala	Ala	Val	Val	Asn	Ala	Phe				
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610		615				620													
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